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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/601,168	07/28/2000	RICHARD BENAROUS	935.38812X00	8585

20457 7590 04/18/2003

ANTONELLI TERRY STOUT AND KRAUS  
SUITE 1800  
1300 NORTH SEVENTEENTH STREET  
ARLINGTON, VA 22209

EXAMINER

SCHNIZER, HOLLY G

ART UNIT	PAPER NUMBER
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1653

DATE MAILED: 04/18/2003

12

Please find below and/or attached an Office communication concerning this application or proceeding.

**FILE COPY**

**Office Action Summary**

Application No.

09/601,168

Applicant(s)

BENAROUS ET AL.

Examiner

Holly Schnizer

Art Unit

1653

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --  
**Period for Reply**

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133).
- Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

**Status**

- 1) ☒ Responsive to communication(s) filed on 15 January 2003.
- 2a) ☒ This action is **FINAL**.                      2b) ☐ This action is non-final.
- 3) ☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

**Disposition of Claims**

- 4) ☒ Claim(s) 1,3,4,6,7 and 22-50 is/are pending in the application.
- 4a) Of the above claim(s) 6,22-30,33-36 and 38-50 is/are withdrawn from consideration.
- 5) ☐ Claim(s) \_\_\_\_\_ is/are allowed.
- 6) ☒ Claim(s) 1,3,4,7,31,32 and 37 is/are rejected.
- 7) ☐ Claim(s) \_\_\_\_\_ is/are objected to.
- 8) ☐ Claim(s) \_\_\_\_\_ are subject to restriction and/or election requirement.

**Application Papers**

- 9) ☐ The specification is objected to by the Examiner.
- 10) ☒ The drawing(s) filed on 29 January 1999 is/are: a) ☒ accepted or b) ☐ objected to by the Examiner.  
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).
- 11) ☐ The proposed drawing correction filed on \_\_\_\_\_ is: a) ☐ approved b) ☐ disapproved by the Examiner.  
If approved, corrected drawings are required in reply to this Office action.
- 12) ☐ The oath or declaration is objected to by the Examiner.

**Priority under 35 U.S.C. §§ 119 and 120**

- 13) ☒ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).  
a) ☒ All b) ☐ Some \* c) ☐ None of:  
1. ☐ Certified copies of the priority documents have been received.  
2. ☐ Certified copies of the priority documents have been received in Application No. \_\_\_\_\_.  
3. ☒ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).  
\* See the attached detailed Office action for a list of the certified copies not received.
- 14) ☐ Acknowledgment is made of a claim for domestic priority under 35 U.S.C. § 119(e) (to a provisional application).  
a) ☐ The translation of the foreign language provisional application has been received.
- 15) ☐ Acknowledgment is made of a claim for domestic priority under 35 U.S.C. §§ 120 and/or 121.

**Attachment(s)**

- 1) ☒ Notice of References Cited (PTO-892)
- 2) ☐ Notice of Draftsperson's Patent Drawing Review (PTO-948)
- 3) ☐ Information Disclosure Statement(s) (PTO-1449) Paper No(s) \_\_\_\_\_
- 4) ☐ Interview Summary (PTO-413) Paper No(s). \_\_\_\_\_
- 5) ☐ Notice of Informal Patent Application (PTO-152)
- 6) ☐ Other: \_\_\_\_\_

**DETAILED ACTION**

***Status of the Claims***

The Amendment filed January 15, 2003 has been entered and considered. Claims 2 and 5 have been cancelled. Therefore, Claims 1, 3-4, 6-7, and 22-50 are pending. Claims 6, 22-26, 27-30, 33-36, and 38-50 are withdrawn from consideration as being drawn to a non-elected invention. Claims 1, 3-4, 7, 31, 32, and 37 will be considered in this Office Action.

**Objections/Rejections Withdrawn**

***Objection to Specification Withdrawn***

The objection to the Specification for recitation of amino acid sequences without reference to a sequence identifier is withdrawn in light of the amendment.

***Claim Objections Withdrawn***

The objection to Claim 1 is withdrawn in light of the amendment.

The objection of Claims 7, 31, and 32 because they encompass polynucleotides encoding peptides devoid of the F box and peptides devoid of WD units and therefore encompass non-elected subject matter is withdrawn in light of the amendment to Claim 7. Correction is required.

***Rejections Withdrawn***

The rejections of Claims 2 and 5 under 35 U.S.C. 112, second paragraph are withdrawn in light of the cancellation of these claims.

The rejection of Claim 27 under 35 U.S.C. 112, second paragraph is withdrawn in light of the amendment making it a non-elected claim.

The rejection of Claim 37 under 35 U.S.C. 112, second paragraph is withdrawn in light of the amendment.

The rejection of Claims 7, 27, 31, 32, and 37 under 35 U.S.C. 112, first paragraph for lack of enablement is withdrawn in light of the amendment removing limitations relating to fragments from the claims.

The rejections of Claims 7, 31, and 32 are rejected under 35 U.S.C. 102(a) and (b) as being anticipated by Skowyra et al., Bour et al., Rubinfeld et al., and Inoue et al. is withdrawn in light of the amendment. Skowyra et al., Bour et al., Rubinfeld et al. and Inoue et al. do not teach or suggest that the proteins taught therein contain 7 WD units as found in SEQ ID NO:2 of the present invention.

**Objections/Rejections**

***Claim Objections***

As stated in the previous Office Action (with reference to original claim 1) the claims should refer to sequence identifiers as "SEQ ID NO:1" as indicated in 37 C.F.R. 1.821(d) rather than as "SEQ ID No. 1" as is presently claimed in Claim 7. Correction is required.

***Claim Rejections - 35 USC § 112***

The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

Claims 1, 3, 4, 7, 31, 32, and 37 are rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention.

Claim 1 is rejected because it is unclear as to whether the claimed protein must only comprise "units having the following positions in SEQ ID NO:2", interpreted to mean that the protein has an F box and 7 WD units, or whether the claimed protein must comprise the sequences having the following positions in SEQ ID NO:2, which would be interpreted to mean that the protein would have to have those identical sequences. Therefore, the metes and bounds of the claim are unclear. Claims 3, 4, 7, 31, 32, and 37 are also rejected since they depend from Claim 1 but do not correct its deficiencies. Clarification is required.

Steps b) and c) of Claim 7 are unclear. First, the claim is drawn to a nucleic acid sequence coding for the h- $\beta$ TrCP protein consisting of "b) a DNA sequence which hybridizes under strict conditions with the above sequence" (the above sequence is SEQ ID NO:1 which encodes the h- $\beta$ TrCP protein). This is confusing since a sequence which hybridizes to SEQ ID NO:1 would be complementary to SEQ ID NO:1 and therefore could not encode the h- $\beta$ TrCP. Second, step c) recites that the DNA

sequence "results from the sequences a) and b) above and codes for the human protein h- $\beta$ TrCP". However, sequence a) is the sequence of SEQ ID NO:1 and sequence b) is a sequence which hybridizes to SEQ ID NO:1. Thus, sequences a) and b) are complimentary. The claim is unclear as to what type of sequence "results from" two sequences that are complimentary to each other. Third, it is unclear as to what type of sequence "results from the sequence" of b) and codes for the human protein h- $\beta$ TrCP since these two sequences are complimentary. Claims 31 and 32 are also rejected since they depend from Claim 7 and do not correct its deficiencies. Correction is required.

The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

Claims 1, 3, 4, 7, 31, and 32 are rejected under 35 U.S.C. 112, first paragraph, as containing subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention. Applicant is referred to the interim guidelines on written Guidelines published January 5, 2001 in the Federal Register, Vol. 66, No. 4, p. 1099-1111 (available at [www.uspto.gov](http://www.uspto.gov)) and the Examiner training Materials on Written Description also available at [www.uspto.gov](http://www.uspto.gov).

The addition of Claims 1, 3, and 4 to this rejection is necessitated by amendment:

Claim 1 has been amended to delete the limitation that the protein has the sequence of SEQ ID NO:2 and to add the limitations previously found in Claim 5. Claim 5 was indicated as allowable because it depended from Claim 1 and thus was drawn to a protein having the sequence of SEQ ID NO:2. However, as amended, the claim is no longer limited to the sequence of SEQ ID NO:2 but may have any number of a wide variety of sequences. Claims 3 and 4 depend from Claim 1 and do not further narrow the genus of sequences claimed.

Applicants contend that the amendment to the claims deleting the limitations relating to peptide fragments is sufficient to overcome the rejection. However, the claims are still drawn to a large genus of protein and nucleic acid sequences as discussed below.

The claims, as amended, are considered to be drawn to any protein or nucleic acid sequence that codes for a protein having the activity of interacting with the Vpu protein of HIV-1, the cell protein I $\kappa$ B, the cell protein  $\beta$ -catenin, or the skp1p protein and having an F-box and seven WD units. The specification discloses a single  $\beta$ -TrCP protein of SEQ ID NO:2 and various modifications of SEQ ID NO:2 wherein a single domain is removed (i.e. SEQ ID NO:2 without an F-box or SEQ ID NO:2 without the first WD domain). The specification and claims do not provide any guidance with respect to the relationship between specific amino acids within the sequence and function. For example, there is no guidance as to what effect changing amino acids within the F-box would have on skp1p binding. The Specification indicates that such changes could affect the binding. For example, page 2 of the Specification states "it is not certain that

the function of the homologous proteins will be totally conserved. Moreover, there are numerous examples which show that there are always significant differences between species" (p. 2, lines 4-6). Thus, the Specification acknowledges that even two highly homologous proteins may not have identical function. However, the Specification does not teach what identifying characteristics of the sequence of the protein of the present invention give it its identifying function. The written description requirement may be satisfied through disclosure of function and minimal structure when there is a well-established correlation between structure and function. In contrast, without such a correlation, the capability to recognize or understand the structure from the mere recitation of function and minimal structure is highly unlikely and is little more than a wish for possession (see Fed. Reg. (2001) Vol. 66(4) p. 1110, Col. 2, citation 49 citing Eli Lilly, 43 USPQ2d at 1406)).

In addition, the specification and claims do not provide any structural or functional characteristics to distinguish a human  $\beta$ -TrCP from that of other species or proteins that have different functions. For example, Hatakeyama et al. (Proc. Natl. Acad. Sci. (1999) 96: 3859-3863), Skowrya et al. (Cell (1997) 91: 209-219) disclose proteins that appear to have the function of the protein of the claimed invention (clm7), that of binding skp1p. The specification does not provide guidance as to whether these proteins of similar function and structure to that of the protein of the present invention are considered modified  $\beta$ -TrCP proteins. Similarly, the specification acknowledges that the previously identified slimb protein and  $\beta$ -TrCP of Xenope are homologs of the protein of the present invention (p.1 lines 19-35). In fact, as shown in Spevak et al.



(Mol. Cell Biol. (1993) 13: 4953-4966), *Xenopus*  $\beta$ -TrCP has an Fbox and seven WD units that are identical to those of SEQ ID NO:2 (as defined in Claim 1) except for a single amino acid difference in the F-box and a single amino acid difference in the second WD unit. However, the specification does not provide any guidance as to what effect changing a single amino acid in the F-box or WD units would have on activity or the effect that modifications outside the F-box and WD units would have on activity. Moreover, the specification does not provide any structural or functional characteristics of a "human"  $\beta$ -TrCP protein so as to distinguish it from homologs from other species.

Therefore, the scope of the claims include innumerable structural variants (nucleic acid sequences that hybridize and proteins of varying sequences that have an Fbox and seven WD domains) and the genus is highly variant because a significant number of structural differences between genus members is permitted. There is no description of mutational sites that exist in nature, and there is no description of how the structure of the specific SEQ ID NO: relates to the function of the protein or disease. Thus, the common attributes of the genus are not described. One of skill in the art would conclude that applicant was not in possession of the claimed genus because a description of only one member of such a large genus is not representative of the variants of the genus and is insufficient to support the claims.

### ***Claim Rejections - 35 USC § 102***

The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

Art Unit: 1653

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

Claims 1, 3, 4, 7, 31, and 32 are rejected under 35 U.S.C. 102(b) as being anticipated by Spevak et al. (Mol. Cell Biol. (1993) 13: 4953-4966).

Spevak et al. disclose a protein that has the units, an F-box and 7 WD domains, as described in present Claim 1. The sequences of the 1<sup>st</sup> and 3rd –7<sup>th</sup> WD units of the protein of Spevak et al. are identical to the corresponding sequences in SEQ ID NO:2 (see figure 9, p. 4960 and sequence alignment attached to this Office Action, claimed positions highlighted). The F-Box and 2nd WD unit each have only one amino acid different than the corresponding units of SEQ ID NO:2 (see attached sequence alignment). Thus, the protein of Spevak et al. is considered to comprise the units having the positions in SEQ ID NO:2 described in Claims 1, 3 and 4. The function of a protein (in this case capability of interacting with proteins degradable by proteasome (clm 1), specifically Vpu, I $\kappa$ B or  $\beta$ -catenin (clm 3) or skp1p (clm 4)) is dependent on its sequence. Two proteins of the same sequence will have the same structure and thus the same function. Thus, since the protein of Spevak et al. has the same sequence and structure as that of the claimed protein, it is an inherent property of the protein described in Spevak et al. that it would have the same function. Thus, claims 1, 3 and 4 are anticipated by Spevak et al.

The DNA sequence encoding the protein disclosed in Spevak et al. is highly homologous to SEQ ID NO:1 (82% local similarity; see sequence alignment attached)

and would hybridize to a polynucleotide having SEQ ID NO:1. Spevak et al. teach that the  $\beta$ TrCP was expressed in cdc20 cells using an expression vector containing the nucleic acid molecule encoding  $\beta$ -TrCP and means for expression, and therefore Spevak et al. meet the limitations of Claims 1, 7, 31, and 32.

It is noted that Claim 1 is unclear as to whether the claimed protein must only comprise "units having the following positions in SEQ ID NO:2", interpreted to mean that the protein has an F box and 7 WD units, or whether the claimed protein must comprise the sequences having the following positions in SEQ ID NO:2, which would be interpreted to mean that the protein would have to have those identical sequences. If the latter is intended then amendment to clarify this intention would overcome the above rejection under 35 U.S.C. 102(b) since the protein of Spevak et al. has a single amino acid difference from that of SEQ ID NO:2 in each of the F-box and 1<sup>st</sup> WD unit.

### ***Conclusions***

No Claims are allowable. A thorough search of the prior art did not reveal any teaching or suggestions of a protein having the sequence of SEQ ID NO:2, a nucleic acid molecule encoding the protein of SEQ ID NO: 2 or a nucleic acid molecule having the sequence of SEQ ID NO:1, or a method of identifying anti-HIV-1 antiviral agents using a protein having the sequence of SEQ ID NO:2.


Applicant's amendment necessitated the new ground(s) of rejection presented in this Office action. Accordingly, **THIS ACTION IS MADE FINAL**. See MPEP § 706.07(a). Applicant is reminded of the extension of time policy as set forth in 37 CFR 1.136(a).

A shortened statutory period for reply to this final action is set to expire **THREE MONTHS** from the mailing date of this action. In the event a first reply is filed within **TWO MONTHS** of the mailing date of this final action and the advisory action is not mailed until after the end of the **THREE-MONTH** shortened statutory period, then the shortened statutory period will expire on the date the advisory action is mailed, and any extension fee pursuant to 37 CFR 1.136(a) will be calculated from the mailing date of the advisory action. In no event, however, will the statutory period for reply expire later than **SIX MONTHS** from the date of this final action.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Holly Schnizer whose telephone number is (703) 305-3722. The examiner can normally be reached on Monday through Wednesday from 8 am to 5:30 pm.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Christopher Low can be reached on (703) 308-2923. The fax phone numbers for the organization where this application or proceeding is assigned are (703) 308-4242 for regular communications and (703) 308-4242 for After Final communications.

Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to the receptionist whose telephone number is (703) 308-0196.

  
Holly Schnizer  
April 16, 2003

  
CHRISTOPHER S. F. LOW  
SUPERVISORY PATENT EXAMINER  
TECHNOLOGY CENTER 1800

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OM protein - protein search, using sw model

Run on: February 20, 2003, 09:55:06 ; Search time 22 Seconds  
(without alignments)  
2486.386 Million cell updates/sec

Title: US-09-601-168b-2  
Perfect score: 3034  
Sequence: 1 MPAAEAVLQEKALKEFNSSSE.....PAAQAEPPRSPTTYTISR 569

Scoring table: BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 283224 seqs, 96134422 residues

Total number of hits satisfying chosen parameters: 283224

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database :  
1: PIR:73:1  
2: PIR:73:2  
3: PIR:73:3  
4: PIR:73:4

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

# SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	2397	85.6	518	2 B48088	beta-transducin re
2	1635.5	53.9	701	2 T16607	hypothetical prote
3	690	22.7	506	2 T50211	WD-repeat protein
4	590.5	19.5	605	2 T38932	probable sulfur me
5	545	18.0	640	2 S49932	MER30 protein - ye
6	531.5	17.5	650	2 T46660	sulfur controller
7	520	17.1	579	2 T22703	hypothetical prote
8	519.5	17.1	267	2 S62507	hypothetical trip-a
9	455.5	15.0	1356	2 T18521	beta transducin-11
10	453	14.9	775	2 T45136	hypothetical prote
11	413.5	13.6	1227	2 AE1810	WD repeat protein
12	399	13.2	779	2 S56245	WD-40 repeat prote
13	396.5	13.1	703	2 T43557	cell division cont
14	378.5	12.5	1189	2 A12453	F-box/WD-repeat pr
15	375	12.4	1747	2 AC1842	WD-repeat protein
16	374	12.3	1526	2 AC2239	WD-40 repeat prote
17	373.5	12.3	1258	2 A12155	WD-repeat protein
18	373.5	12.3	1683	2 AE2071	WD-40 repeat prote
19	361.5	11.9	677	2 AE1861	serine/threonine k
20	358.5	11.8	409	2 AB2202	hypothetical prote
21	354	11.7	409	2 S36113	LIS-1 protein - hu
22	354	11.7	410	2 S48052	platelet-activatin
23	341	11.2	515	2 S19487	hypothetical prote
24	339.5	11.2	777	2 T41075	hypothetical prote
25	337	11.1	777	2 A55532	myosin-heavy-chain
26	336	11.1	1146	2 AH2195	hypothetical prote
27	334	11.0	589	2 AG2400	WD-repeat protein
28	333.5	11.0	1711	2 AD1842	WD-40 repeat prote
29	332.5	11.0	317	2 T46032	WD-40 repeat regul

30	330	10.9	333	2 G85034	probable WD-repeat
31	330	10.5	934	2 AG1889	WD-40 repeat prote
32	318	10.5	422	2 A56640	CDc4 repeat unit-c
33	313.5	10.3	376	2 T13266	hypothetical prote
34	312.5	10.3	714	2 S56893	hypothetical prote
35	312	10.3	786	2 AG2375	WD-40 repeat-prote
36	312	10.3	876	2 T51507	WD-repeat protei
37	307.5	10.1	1693	2 S76086	beta transducin-11
38	306.5	10.1	598	2 AE2415	WD-repeat protein
39	306	10.1	1189	2 AH2154	transcription init
40	305	10.1	704	2 S33263	WD-repeat protein
41	299.5	9.9	1194	2 T03818	apoptotic proteina
42	298	9.8	265	2 AF1890	WD-repeat protein
43	297.5	9.8	357	2 A12099	WD-40 repeat prote
44	297	9.8	502	2 T41148	tip-as repeat con
45	296.5	9.8	304	2 AG1837	WD-40 repeat prote

## ALIGNMENTS

### RESULT 1

B48088  
beta-transducin repeat-containing protein - African clawed frog

N:Alternate names: beta-Trop  
C:Species: Xenopus laevis (African clawed frog)

C:Date: 26-May-1994 #sequence\_revision 26-May-1994 #text\_change 21-Jul-2000

C:Accession: B48088

R:Spencer, M., Kasper, B.D., Stralowa, C., Castanon, M.J.  
Mol. Cell. Biol. 13, 4953-4966, 1993

A:Title: Saccharomyces cerevisiae cdc4 mutants arrested at a late stage in anaphase

A:Reference number: A48088; MUID:3330289; PMID:8393141

A:Accession: B48088

A:Molecule type: mRNA

A:Residues: 1-518 <SPE>

A:Cross-references: GB:98268; NID:q295542; PIDN:AA02810.1; PID:q295543

C:Superfamily: unassigned WD repeat proteins; WD repeat homology

C:Keywords: duplication

F:431-462/Domain: WD repeat homology <MD1>

Query Match	Best Local Similarity	Score	DB 2;	Length	518;
Matches 488; Conservative	91.6%;	Pred. No. 1.2e-193;	Mismatches 8;	Indels 30;	Gaps 1;
QY 18	SSREDCNNGEPKRIIPKNSLRQTNSCARCLQNOETVCLASTAKTENCVAKTLAN 77				
DB 13	ASEREDCNRPDPKRIIPKNTLRQ-----TKLAN 42				
QY 78	GTSSMIVPKQKRLASYEKEKELCVYKFEQMSDQVEFEVHLISQCHQHGHSYK 137				
DB 43	GTSSMIVPKQKRLASANEKEKELCVYKFEQMSDQVEFEVHLISQCHQHGHSYK 102				
QY 138	PMQORDITVLPARGDHTAENTISTIDAKSICMAHVCENYVSDGMNGLTERAV 197				
DB 103	PMQORDITVLPARGDHTAENTISTIDAKSICMAHVCENYVSDGMNGLTERAV 162				
QY 198	RTDSLWGLAERGMGOYLKPKPDGNAPPNSFYALPKTIQDLETESNWRGSHSL 257				
DB 163	RTDSLWGLAERGMGOYLKPKPDGNAPPNSFYALPKTIQDLETESNWRGSHSL 222				
QY 258	QRTCHSESTSKGYTCIQYDQKIVSGIRNTLTKIQKNTLECKRIILGHVSGVZLEQYD 317				
DB 223	QRTCHSESTSKGYTCIQYDQKIVSGIRNTLTKIQKNTLECKRIILGHVSGVZLEQYD 282				
QY 318	RVITGSSDSTAVYVWVNTNGEMANTGHISEVYHREYNGNMAVYCSKDRSTAVYMASP 377				
DB 283	RVITGSSDSTAVYVWVNTNGEMANTGHISEVYHREYNGNMAVYCSKDRSTAVYMASP 342				
QY 378	TITTLRRVLGHRRAVNVVDFOCKIVASGDRITKVTSTCEVVRILAGKRGICLQ 437				
DB 343	TITTLRRVLGHRRAVNVVDFOCKIVASGDRITKVTSTCEVVRILAGKRGICLQ 402				

OY 438 /RDLVWSSGSSDNTIRLMDIEGACLRVLGSEHEWCAQNDKRYSGAYDCKIKYMDL 497  
 DB 403 YRDLVWSSGSSDNTIRLMDIEGACLRVLGSEHEWCAQNDKRYSGAYDCKIKYMDL 462  
 OY 498 VVALDPPAPAGTICLRLLGSGRVEFLDDEFOIYSSSHDITILMDPLNDP 550  
 DB 463 VVALDPPAPAGTICLRLLGSGRVEFLDDEFOIYSSSHDITILMDPLNDP 515

## RESULT 2

T16607

hypothetical protein K10B2.1 - *Caenorhabditis elegans*  
 C:Species: *Caenorhabditis elegans*  
 C:Date: 20-Sep-1999 #sequence\_revision 20-Sep-1999 #text\_change 20-Sep-1999  
 C:Accession: T16607  
 R:Miller, N.

submitted to the EMBL Data Library, June 1995  
 A:Description: The sequence of C. elegans, cosmid K10B2.  
 A:Accession number: Z16545  
 A:Reference number: T16607

A:Status: preliminary; translated from GB/EMBL/DBJ  
 A:Molecule type: DNA  
 A:Residues: 1-701 <MIL>

A:Cross-references: EMBL:U28730; NID:9860694; PID:9860695; PIDN:AAA68258.1; CESP:K10B2.1  
 A:Experimental source: strain Bristol N2  
 A:Genetics:  
 A:gene: CESP:K10B2.1  
 A:introns: 78/3; 125/1; 183/2; 281/3; 404/3; 551/3; 668/3

Query Match 53.9%; Score 1635.5; DB 2; Length 701;  
 Best Local Similarity 57.7%; Pred. No. 8.1e-119;  
 Matches 328; Conservative 69; Mismatches 116; Indels 55; Gaps 8;

OY 31 RKIIPEKNSIRQTYNSCARLCLNQTVCYLAETAMKTECNCAKTKLAN-----GNS 80  
 DB 2 RRRREKRAKMGKRRARDGSGIALVVCST-----IERCF--TAVSNPIFFLESTFFSVF 55  
 OY 81 SMIVPQR-----KISATYERKEKLCVKYFEQNSQSDQVEFEHLISQCHQHGHSINSY 135  
 DB 56 SFLEPSRNQIQLSRSFSSEFVL-----KWEHEQLDPMKIVHRLSHVQLGKVNMF 110

OY 136 LKPNLORDFTLPAAGLDHIAENLSTYDASLCAELVCKEYTVTSDGMKMLIER 195  
 DB 111 IRPMLORDTSLNPA---HLVELLFINVNSDLKSCSEVTSWRCALARGQHWKMLIK 166  
 OY 196 MVRTDSLWGLAERRGMYL-----FKNRPDGNAPNSFYAL 235  
 DB 167 NVASDSLWGLSEKRWMDKFINISRDMASRYRICEKNYDVNIKRLDOLILMAHYFVSKL 226

OY 236 YRRIIDDIETISNRCGHSIQRHCRSETSKGYCLOYDOKIYSGLRDNTIKIMDN 295  
 DB 227 YRRIIDHININMNRKGYKMTIRINCOSENSGYCLOYDOKIYSGLRDNTIKIMDKR 286  
 OY 296 TLECKRIHGTGYSVLCLOYDERVITTSSTSVYRWVDVNTGEMTLTILHCEAVHLRF 355  
 DB 287 DYSCHSIIISGHGYSVLCLOYDNRVITSSGSDATVRWVDVETGCITLTHCEAVHLRF 346

OY 356 NNGMWTGSKDSIAVMDASPTDITLRVLYGHRVAVVDVDDKYIYASAGDRTIKYW 415  
 DB 347 ANGIWYTSKDSIAVMDVSPDITIRVLYGHRVAVVDVDDKYIYASAGDRTIKYW 406  
 OY 416 NNTSGEYVTLNGHKGACIACLOYRDLVYSGSSDNTIRLMDIEGACLRVLGSEHELVRC 475  
 DB 407 SHDTLEFVTLNGHKGACIACLOYRDLVYSGSSDNTIRLMDIEGACLRVLGSEHELVRC 466

OY 476 IRFDNRRIYSGAYDCKIKYMDLVVALDPPAPAGTICLRLLGSGRVEFLDDEFOIYSS 535  
 DB 467 IRFDNRRIYSGAYDCKIKYMDLVVALDPPAPAGTICLRLLGSGRVEFLDDEFOIYSS 526  
 OY 536 SHDITILMDPLNDPAAQEPSPSRT 563  
 DB 527 SHDITILMDPLNDPAAQEPSPSRT 549

## RESULT 3

T50211

WD-repeat protein [imported] - fission yeast (*Schizosaccharomyces pombe*)  
 C:Species: *Schizosaccharomyces pombe*  
 C:Date: 09-Jun-2000 #sequence\_revision 09-Jun-2000 #text\_change 02-Sep-2000  
 C:Accession: T50211  
 R:McDougal, R.C.; Rajandream, M.A.; Barrell, B.G.; Brown, S.; Murphy, L.; Jones, L.;  
 submitted to the EMBL Data Library, January 2000

A:Accession number: T50211  
 A:Reference number: Z25046  
 A:Status: preliminary; translated from GB/EMBL/DBJ  
 A:Molecule type: DNA  
 A:Residues: 1-506 <MCD>

A:Cross-references: EMBL:AL136538; PIDN:CA866464.1; GSPDB:GN00066; SPDB:SPAC30.05  
 A:Experimental source: strain 972h(-); cosmid G30  
 A:Genetics:  
 A:gene: SPAC296.01; SPDB:SPAC30.05  
 A:Map position: 1  
 A:introns: 43/1; 74/3  
 C:Superfamily: unassigned WD repeat proteins; WD repeat homology

Query Match 22.7%; Score 690; DB 2; Length 506;  
 Best Local Similarity 30.4%; Pred. No. 1.3e-45;  
 Matches 163; Conservative 104; Mismatches 167; Indels 102; Gaps 15;

OY 67 ENCVAK-----TKLNGTSSMIVPQRKISATYERKEKLCVKYFEQNSQSDQVEFEHLIS 122  
 DB 8 KNVKSVSDLTSCSDFSTSSPCLNPLS-----HENNRIDILRDLA 50

OY 123 QMCHYOHGHSINSYLPMDIORDITLPAAGLDHIAENLSTYDASLCAELVCKEYTVTSDGMKMLIER 182  
 DB 51 SLKRGVAVVYNHRSKSLFTFTEVP-----EVLAVFTSLDLDLCKKMLSKMKRL 106

OY 183 TSDGMKMLI-----ERVVRTDSLWGLA-----LAERRGNG----- 213  
 DB 107 LEDPGIMKALYKMGKRRARDGSGIALVVCST-----IERCF--TAVSNPIFFLESTFFSVF 55

OY 214 OYLFKNRPDGNAPNSFYALPKIIDDITISNRCGHSIQRHCRSETSKGYCLOYDOKIYSGLRDNTIKIMDN 295  
 DB 167 QPFI-----DSNGRPLWMSYLV---KEHAHLDNMRHGFVSTPNNSIREPADODF 217

OY 265 -ETSKGYCLOYDOKIYSGLRDNTIKIMDNTECKRIITLGHGYSVLCLOYDER-VII 321  
 DB 218 RATLDSVYCVQYDEIWSGSKDRTVSWVDNYSKFLIKLKGHSVLCIDFCRRRLV 277

OY 322 TGSSDSIVRWVDVNTGEMTLTILHCEAVHLRFNNGMWTGSKDSIAVW-DMASPTD 379  
 DB 278 SGSSDSTIIMDMONRRPLKYFGHTDNLGVVSENYIISSRDHTARVRLDATSPE 337

OY 380 ITRRLVLYGHRVAVVDVDDKYIYASAGDRTIKIMDNSTGCFVTLNGHKGACIAC 437  
 DB 338 ACM-HVLRGLASVNSVOYSKTLIVTASDSRLRTKWDITTGHCRTIHAHORGIAACA 396

OY 438 YRDLVWSSGSSDNTIRLMDIEGACLRVLGSEHEWCAQNDKRYSGAYDCKIKYMDL 497  
 DB 397 YNGKFIVSGSSDLTIRLEASSGKLMLOGHDLITVAFNDEKTIYSGDYDQVIAVW- 455

OY 498 VVALDPPAPAGTICLRLLGSGRVEFLDDEFOIYSSSHDITILMDPLNDP 550  
 DB 456 -----FNTGEQHCVLHNSRNSRVPLGDFHRRRIIACVSHSEILWNF 497

RESULT 4  
 T38932  
 Probable sulfur metabolite control protein - fission yeast (*Schizosaccharomyces pombe*)  
 C:Species: *Schizosaccharomyces pombe*  
 C:Date: 03-Dec-1999 #sequence\_revision 03-Dec-1999 #text\_change 26-May-2000  
 C:Accession: T38932  
 R:Badcock, K.; Churcher, C.M.; Wood, V.; Barrell, B.G.; Rajandream, M.A.;  
 submitted to the EMBL Data Library, April 1997  
 A:Reference number: Z21818  
 A:Accession: T38932



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||||| 1141 ACCCTGAGGAGGCTGCTGGGAGACCCGAGCTGCGTCAATGTTGATGATGAC 1200
||||| 1270 AAGTACATTTTCTGATGATGAGATAGAACTATTAAGATATGAACAAAGTACTTCT 1329
||||| 1201 AAGTACATGCTTTCTGCTCTGAGATAGACACATTAAGTGTGGAACAAAGTACTTCT 1260
||||| 1330 GAATTTGTAAGACCTTAATGAAGACAAAGAGGATTCCTGTTTGGAGTACAGAGAC 1389
||||| 1261 GAATTTGTAAGACCTTAATGAAGACAAAGGATTCCTGTTTGGAGTACAGAGAC 1320
||||| 1390 AGGCTGAGTGTGAGTGTGATGATGATGATGATGATGATGATGATGATGATGATGAT 1449
||||| 1321 AGGCTGAGTGTGAGTGTGATGATGATGATGATGATGATGATGATGATGATGATGAT 1380
||||| 1450 GCATGTTTACGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGAT 1509
||||| 1381 GCATGTTTACGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGAT 1440
||||| 1570 TTGAGACCCCGCTGCTGCTGAGGACATCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 1629
||||| 1501 TTGAGACCCCGCTGCTGCTGAGGACATCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 1560
||||| 1630 AGAGTTTTCGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGAT 1689
||||| 1561 AGAGTTTTCGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGAT 1620
||||| 1690 ATCCGATGCTGAGTCTCTCAATGATGATGATGATGATGATGATGATGATGATGATGAT 1749
||||| 1621 ATCTGATGCTGAGTCTCTCAATGATGATGATGATGATGATGATGATGATGATGATGAT 1680
||||| 1750 TCTGCAACATACACATCTTCCAGATTA 1779
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RESULT 12  
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LOCUS African clawed frog beta-TrCP mRNA, complete cds.  
ACCESSION M98268  
VERSION M98268.1 GI:295542  
KEYWORDS beta-transducin repeats.  
Xenopus laevis (library: S. cerevisiae expression library of  
X.laevis oocytes) cDNA to mRNA.  
Xenopus laevis  
ORIGIN Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Amphibia; Batrachia; Anura; Mesobatrachia; Pipridae; Pipidae;  
Xenopodidae; Xenopus.  
1 (bases 1 to 1671)  
SPEAK, M., Keiper, B.D., Stratowa, C. and Castanon, M.J.  
Saccharomyces cerevisiae cdc15 mutants arrested at a late stage in  
anaphase are rescued by Xenopus CDNs encoding N-ras or a protein  
with beta-transducin repeats  
MOL. CELL. BIOL. 13 (8): 4953-4966 (1993)  
JOURNAL 93370289  
MEDLINE 8393141  
PUBMED  
FEATURES  
Location/Qualifiers  
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GHRGIACLOYRDLVYSSSDNTRIMDEGCACLRVLEGHEILVRCIFRDKRIVS
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1671
polya_site
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BASE COUNT 495 a 341 c 424 g 411 t
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Query Match 47.3%; Score 1017.8; DB 5; Length 1671;
Best Local Similarity 81.8%; Pred. No. 1e-278;
Matches 1175; Conservative 0; Mismatches 262; Indels 0; Gaps 0;

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344 CAAGCTATGAAAGAAAGAAAGAACTGTGTGCAAAATCTTTGAGCAGTGGTCAAGTCA 403  
217 CAATTTGCAAGAAAGAAAGAAAGAACTGTGTGCAAAATCTTTGAGCAGTGGTCAAGTCA 276  
404 ATCAAGTGAATTTGTGGAACATCTTATATCCCAATATGTCTATTACCAACATGGGCACA 463  
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464 TAACTGTATCTTAAACCTATGTTGAGAGAGATTTCAATACCTGCTCTCGAGCTGGG 523  
337 TAAACACTTACCTTAAACCTATGTTGAGAGAGATTTCAATACCTGCTCTCGAGCTGGG 396  
524 GATTGATCATATGCTGAGAAACATCTGTCTATACCTGATGCAAAATCTATGTGCTG 583  
397 GACTGATCATATGCTGAGAAACATCTGTCTATACCTGATGCAAAATCTATGTGCTG 456  
584 CTGAACCTGTGTGAGAAACATCTGTCTATACCTGATGCAAAATCTATGTGCTG 643  
457 CAGAACCTGTGTGAGAAACATCTGTCTATACCTGATGCAAAATCTATGTGCTG 516  
644 TTATCGAGAGAAATGCTGAGAGAGATTTCTGTGAGAGAGCTGGCAGAAAGAGAT 703  
517 TCATAGAGCGGATGCTGAGAGAGATTTCTGTGAGAGAGCTGGCAGAAAGAGAT 576  
704 GGGGACAGTATTTATTTAAACAAACCTCTGAGGAAATGCTCTCCCACTCTTTT 763  
577 GGGGACAGTATTTATTTAAACAAACCTCTGAGGAAATGCTCTCCCACTCTTTT 636  
764 ATAGAGACCTTTATTTAAACAAACCTCTGAGGAAATGCTCTCCCACTCTTTT 823  
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817 GGGTAAACAAACATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGAT 876  
1004 GTTCCAGTATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGAT 1063  
877 GTTCCAGTATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGATGAT 936



1630 AGAGTTTTCAGTACAGTTTGTATGATTCAGATTGTGTCAGTATTCACATGATGACACA 1689  
 1561 AGAGTTTTCAGTACAGTTTGTATGATTCAGATTGTGTCAGTATTCACATGATGACACA 1620  
 1621 ATTCTCATCTGGAGTCTCTGTAATGATCAGAGTGTCTCAGACCGTGAACCCCGCTCCCT 1749  
 1750 TCTGACATACACTACATCTCCAGATTAAT 1781  
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RESULT 11  
 AF112979 1710 bp mRNA linear ROD 02-MAR-1999  
 LOCUS Mus musculus beta-transducin repeat containing protein mRNA,  
 DEFINITION complete cds.  
 ACCESSION AF112979  
 VERSION AF112979.1 GI:4140717  
 KEYWORDS

SOURCE Mus musculus.

ORGANISM

Mus musculus.  
 Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.

REFERENCE 1 (bases 1 to 1710)  
 Spencer, E., Jiang, J. and Chen, Z. J.

Signal-induced ubiquitination of IkappaBalpha by the F-box protein

Slmb/beta-trcp

Genes Dev. 13 (3), 284-294 (1999)

JOURNAL MEDLINE 99145465

PUBMED 990853

REFERENCE 2 (bases 1 to 1710)  
 Chen, Z. J.

Direct Submission

Submitted (10-DEC-1998) Molecular Biology and Oncology, UT

Southwestern Medical Center, 5323 Harry Hines Blvd, Dallas, TX

75235-9148, USA

FEATURES

source

location/Qualifiers

1..1710

/organism="Mus musculus"

/db\_xref="taxon:10090"

1..1710

/function="ubiquitinates phosphorylated IkB"

/note="beta-trcp, F-box protein; IkB-ubiquitin ligase;  
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 Homo sapiens beta-trcp and Drosophila melanogaster Slmb"

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 ISR"

BASE COUNT 469 a 399 c 453 g 389 t

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Query Match 67.1%; Score 1442.8; DB 10; Length 1710;  
 Best Local Similarity 90.2%; Pred. No. 0;  
 Matches 1543; Conservative 0; Mismatches 167; Indels 0; Gaps 0;

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DB 1 ATGACCCGGGCGAGGCGTGTGCAAGAGAGGCACTAAGTTATGATTCCTCAGAG 60

QY 130 AGAGACAGTGTATATATATGCGAACCCTCAGGAGATATATACACAGAAAGATTCATT 189

DB 61 AGAGAAACAGTATATATATATGCGAACCCTCAGGAGATATATACACAGAAAGATTCATT 120

QY 190 AGACAGCATACACAGAGCTGTGCGAGACTCTGCTTAACCAAGAAACAGATATTAGCA 249

DB 121 AGACAGCATACACAGAGCTGTGCGAGACTCTGCTTAACCAAGAAACAGATATTAGCA 180

QY 250 AGACAGCTGTATGAAAGCTGGAATTTGTGTGCGCAAAACAAACTTCCCAATGCGACTTC 309

DB 181 AGACAGCTGTATGAAAGCTGGAATTTGTGTGCGCAAAACAAACTTCCCAATGCGACTTC 240

QY 310 AGTATGTTGTGCGCAAGCAAGCAAACTCTCAGCAAGCTATGAAAGAAAGCAAGCA 369

DB 241 AGCATGTTGTGCGCAAGCAAGCAAACTCTCAGCAAGCTATGAAAGAAAGCAAGCA 300

QY 370 TGTGTCAATACCTTATGAGAGTGTGCGAGAGTCAAGATCAAGTGAATTTGTGCAATCTT 429

DB 301 TGTGTCAATACCTTATGAGAGTGTGCGAGAGTCAAGATCAAGTGAATTTGTGCAATCTT 360

QY 430 ATATCCCAATGTGTATACCAACATGGGCAATTAATCTGTATCTTAACCTATGTTG 489

DB 361 ATATCCCAATGTGTATACCAACATGGGCAATTAATCTGTATCTTAACCTATGTTG 420

QY 490 CAGAGATTTTCAATACCTCTGCGAGCTCGGGAGTTGATCATATGCTGAGAACTT 549

DB 421 CAGAGATTTTCAATACCTCTGCGAGCTCGGGAGTTGATCATATGCTGAGAACTT 480

QY 550 CTGTACCTGAGTCCCAATCACTATGCTGCTGAACCTTGTGTGCAAGAAAGTATC 609

DB 481 CTGTACCTGAGTCCCAATCACTATGCTGCTGAACCTTGTGTGCAAGAAAGTATC 540

QY 610 CGAGTACCTGAGTCCCAATCACTATGCTGCTGAACCTTGTGTGCAAGAAAGTATC 669

DB 541 CGAGTACCTGAGTCCCAATCACTATGCTGCTGAACCTTGTGTGCAAGAAAGTATC 600

QY 670 TCTGTGTGAGAGGCGCTGTGCGAGAGAGAGTGGGAGTATTTATTCAAAACAA 729

DB 601 TCTGTGTGAGAGGCGCTGTGCGAGAGAGAGTGGGAGTATTTATTCAAAACAA 660

QY 730 CCTCTGAGGGAATGCTCTCCCACTTTTATAGACACTTATTCATAATATTA 789

DB 661 CCTCTGAGGGAATGCTCTCCCACTTTTATAGACACTTATTCATAATATTA 720

QY 790 CAAGACATGAGACATATGATCTAATGAGAGTGTGGAAGACATAGTTTACAGAAAT 849

DB 721 CAAGACATGAGACATATGATCTAATGAGAGTGTGGAAGACATAGTTTACAGAAAT 780

QY 850 CACTGCCGAGTGAACCAAGCAAGAGTTCCTGTTTACAGTATGATGACAGAAATA 909

DB 781 CACTGCCGAGTGAACCAAGCAAGAGTTCCTGTTTACAGTATGATGACAGAAATA 840

QY 910 GTAAGGCGCTTCCGAGACACACATCAAGTCTGGATTAACCAATGGAATGCAAG 969

DB 841 GTAAGGCGCTTCCGAGACACACATCAAGTCTGGATTAACCAATGGAATGCAAG 900

QY 970 CGAATTCACAGGCGATACAGTTCAGTCTGCTGCTCCAGTATGATGAGAGTATC 1029

DB 901 CGAATTCACAGGCGATACAGTTCAGTCTGCTGCTCCAGTATGATGAGAGTATC 960

QY 1030 ATACAGATATATCGATTCAGGTCAGAGTGTGAGTGAATTAACAGTGAATCTTA 1089

DB 961 ATACAGATATATCGATTCAGGTCAGAGTGTGAGTGAATTAACAGTGAATCTTA 1020

QY 1090 AACAGGTCATTCACATGTCAGAGCACTTGTGCACTTGTGCAATTAATGCGATATG 1149

DB 1021 AACAGGTCATTCACATGTCAGAGCACTTGTGCACTTGTGCAATTAATGCGATATG 1080

QY 1150 GTGACCTGCTCAAGATATGCTTCATGCTGTATGAGTATGCGCTCCCAATGACATT 1209

DB 1081 GTGACCTGCTCAAGATATGCTTCATGCTGTATGAGTATGCGCTCCCAATGACATT 1140

QY 1210 ACCCTCCGAGGAGTGTGTGTGAGACACCGAGCTGCTCATATGTTGATGATGAC 1269